

DOCKET NO.: CARP-0057

PATENT

APPENDIX C

| Claim Limitation | Support in 1989 GB Application |
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| 28. A humanized immunoglobulin having complementarity determining regions (CDRs) from a donor immunoglobulin and heavy and light chain variable region frameworks from human acceptor immunoglobulin heavy and light chains | See page 1, lines 1-2 and 10-20; page 5, lines 8, to page 6, line 4; , and page 8. |
| which humanized immunoglobulin specifically binds to an antigen with an effective antigen binding affinity | See page 5, lines 1-7; page 22, lines 27-35, page 23, lines 5-9, page 24, lines 1-4; page 25, lines 27-33; page 26 last paragraph. |
| wherein said humanized immunoglobulin comprises amino acids from the donor immunoglobulin framework outside both the Kabat CDRs and the structural loop CDRs of the variable regions, | See page 5, lines 1-7; page 26, last paragraph, to page 27, top paragraph. |
| wherein the donor amino acids replace corresponding amino acids in the acceptor immunoglobulin heavy or light chain frameworks, | See page 5, line 8, to page 6, line 4; page 7, lines 5-20. |
| and each of said donor amino acids contributes to antigen binding as determined by X-ray crystallography. | Page 18, lines 11-17, and lines 33-37, and Figs. 20-21 of the application as filed reference residues that may "contribute to antigen binding" as determined using X-ray crystallography. Residues 48, 49, 71, 73, 76, 78, 88, and 91 are identified in Figure 21. |